

SEQUENCE LISTING

<110> Abbott Laboratories
Davidson, Donald J.

<120> NOVEL ANTIANGIOGENIC PEPTIDES,
POLYNUCLEOTIDES ENCODING SAME AND METHODS FOR INHIBITING
ANGIOGENESIS

<130> 5940.US.P3

<140> 08/924,287

<141> 1997-09-05

<150> US 08/851,350

<151> 1997-05-05

<150> US 08/832,087

<151> 1997-04-03

<150> US 08/643,219

<151> 1996-05-03

<160> 40

<170> FastSEQ for Windows Version 4.0

<210> 1

<211> 791

<212> PRT

<213> Homo sapiens

<400> 1

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Val	Thr	Lys	Lys	Gln	Leu	Gly	Ala	Gly	Ser	Ile	Glu	Glu	Cys	Ala	Ala
			20					25					30		
Lys	Cys	Glu	Glu	Asp	Glu	Glu	Phe	Thr	Cys	Arg	Ala	Phe	Gln	Tyr	His
		35					40					45			
Ser	Lys	Glu	Gln	Gln	Cys	Val	Ile	Met	Ala	Glu	Asn	Arg	Lys	Ser	Ser
	50					55					60				
Ile	Ile	Ile	Arg	Met	Arg	Asp	Val	Val	Leu	Phe	Glu	Lys	Lys	Val	Tyr
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Leu	Ser	Glu	Cys	Lys	Thr	Gly	Asn	Gly	Lys	Asn	Tyr	Arg	Gly	Thr	Met
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Ser	Lys	Thr	Lys	Asn	Gly	Ile	Thr	Cys	Gln	Lys	Trp	Ser	Ser	Thr	Ser
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Pro	His	Arg	Pro	Arg	Phe	Ser	Pro	Ala	Thr	His	Pro	Ser	Glu	Gly	Leu
		115					120					125			
Glu	Glu	Asn	Tyr	Cys	Arg	Asn	Pro	Asp	Asn	Asp	Pro	Gln	Gly	Pro	Trp
	130					135					140				
Cys	Tyr	Thr	Thr	Asp	Pro	Glu	Lys	Arg	Tyr	Asp	Tyr	Cys	Asp	Ile	Leu
145					150					155				160	
Glu	Cys	Glu	Glu	Glu	Cys	Met	His	Cys	Ser	Gly	Glu	Asn	Tyr	Asp	Gly
				165					170					175	
Lys	Ile	Ser	Lys	Thr	Met	Ser	Gly	Leu	Glu	Cys	Gln	Ala	Trp	Asp	Ser
			180					185						190	

Gln	Ser	Pro	His	Ala	His	Gly	Tyr	Ile	Pro	Ser	Lys	Phe	Pro	Asn	Lys
		195					200					205			
Asn	Leu	Lys	Lys	Asn	Tyr	Cys	Arg	Asn	Pro	Asp	Arg	Glu	Leu	Arg	Pro
	210					215					220				
Trp	Cys	Phe	Thr	Thr	Asp	Pro	Asn	Lys	Arg	Trp	Glu	Leu	Cys	Asp	Ile
225					230					235					240
Pro	Arg	Cys	Thr	Thr	Pro	Pro	Pro	Ser	Ser	Gly	Pro	Thr	Tyr	Gln	Cys
			245					250						255	
Leu	Lys	Gly	Thr	Gly	Glu	Asn	Tyr	Arg	Gly	Asn	Val	Ala	Val	Thr	Val
			260					265						270	
Ser	Gly	His	Thr	Cys	Gln	His	Trp	Ser	Ala	Gln	Thr	Pro	His	Thr	His
		275					280					285			
Asn	Arg	Thr	Pro	Glu	Asn	Phe	Pro	Cys	Lys	Asn	Leu	Asp	Glu	Asn	Tyr
	290					295					300				
Cys	Arg	Asn	Pro	Asp	Gly	Lys	Arg	Ala	Pro	Trp	Cys	His	Thr	Thr	Asn
305					310					315					320
Ser	Gln	Val	Arg	Trp	Glu	Tyr	Cys	Lys	Ile	Pro	Ser	Cys	Asp	Ser	Ser
			325						330					335	
Pro	Val	Ser	Thr	Glu	Gln	Leu	Ala	Pro	Thr	Ala	Pro	Pro	Glu	Leu	Thr
			340					345					350		
Pro	Val	Val	Gln	Asp	Cys	Tyr	His	Gly	Asp	Gly	Gln	Ser	Tyr	Arg	Gly
		355					360					365			
Thr	Ser	Ser	Thr	Thr	Thr	Thr	Gly	Lys	Lys	Cys	Gln	Ser	Trp	Ser	Ser
	370					375					380				
Met	Thr	Pro	His	Arg	His	Gln	Lys	Thr	Pro	Glu	Asn	Tyr	Pro	Asn	Ala
385					390					395					400
Gly	Leu	Thr	Met	Asn	Tyr	Cys	Arg	Asn	Pro	Asp	Ala	Asp	Lys	Gly	Pro
			405						410					415	
Trp	Cys	Phe	Thr	Thr	Asp	Pro	Ser	Val	Arg	Trp	Glu	Tyr	Cys	Asn	Leu
			420					425					430		
Lys	Lys	Cys	Ser	Gly	Thr	Glu	Ala	Ser	Val	Val	Ala	Pro	Pro	Pro	Val
		435					440				445				
Val	Leu	Leu	Pro	Asp	Val	Glu	Thr	Pro	Ser	Glu	Glu	Asp	Cys	Met	Phe
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Gly	Asn	Gly	Lys	Gly	Tyr	Arg	Gly	Lys	Arg	Ala	Thr	Thr	Val	Thr	Gly
465					470					475					480
Thr	Pro	Cys	Gln	Asp	Trp	Ala	Ala	Gln	Glu	Pro	His	Arg	His	Ser	Ile
			485						490					495	
Phe	Thr	Pro	Glu	Thr	Asn	Pro	Arg	Ala	Gly	Leu	Glu	Lys	Asn	Tyr	Cys
			500					505					510		
Arg	Asn	Pro	Asp	Gly	Asp	Val	Gly	Gly	Pro	Trp	Cys	Tyr	Thr	Thr	Asn
		515					520					525			
Pro	Arg	Lys	Leu	Tyr	Asp	Tyr	Cys	Asp	Val	Pro	Gln	Cys	Ala	Ala	Pro
	530					535					540				
Ser	Phe	Asp	Cys	Gly	Lys	Pro	Gln	Val	Glu	Pro	Lys	Lys	Cys	Pro	Gly
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Arg	Val	Val	Gly	Gly	Cys	Val	Ala	His	Pro	His	Ser	Trp	Pro	Trp	Gln
			565						570					575	
Val	Ser	Leu	Arg	Thr	Arg	Phe	Gly	Met	His	Phe	Cys	Gly	Gly	Thr	Leu
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Ile	Ser	Pro	Glu	Trp	Val	Leu	Thr	Ala	Ala	His	Cys	Leu	Glu	Lys	Ser
		595					600					605			
Pro	Arg	Pro	Ser	Ser	Tyr	Lys	Val	Ile	Leu	Gly	Ala	His	Gln	Glu	Val
	610					615					620				
Asn	Leu	Glu	Pro	His	Val	Gln	Glu	Ile	Glu	Val	Ser	Arg	Leu	Phe	Leu
625					630					635					640
Glu	Pro	Thr	Arg	Lys	Asp	Ile	Ala	Leu	Leu	Lys	Leu	Ser	Ser	Pro	Ala
				645					650					655	

Val Ile Thr Asp Lys Val Ile Pro Ala Cys Leu Pro Ser Pro Asn Tyr
 660 665 670
 Val Val Ala Asp Arg Thr Glu Cys Phe Ile Thr Gly Trp Gly Glu Thr
 675 680 685
 Gln Gly Thr Phe Gly Ala Gly Leu Leu Lys Glu Ala Gln Leu Pro Val
 690 695 700
 Ile Glu Asn Lys Val Cys Asn Arg Tyr Glu Phe Leu Asn Gly Arg Val
 705 710 715 720
 Gln Ser Thr Glu Leu Cys Ala Gly His Leu Ala Gly Gly Thr Asp Ser
 725 730 735
 Cys Gln Gly Asp Ser Gly Gly Pro Leu Val Cys Phe Glu Lys Asp Lys
 740 745 750
 Tyr Ile Leu Gln Gly Val Thr Ser Trp Gly Leu Gly Cys Ala Arg Pro
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 Asn Lys Pro Gly Val Tyr Val Arg Val Ser Arg Phe Val Thr Trp Ile
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 Glu Gly Val Met Arg Asn Asn
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<210> 2
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 <213> Artificial Sequence

<220>
 <223> PCR Amplification Primer

<400> 2
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 45

<210> 3
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 <212> DNA
 <213> Artificial Sequence

<220>
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<400> 3
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<210> 4
 <211> 40
 <212> DNA
 <213> Artificial Sequence

<220>
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<400> 4
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<210> 5
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 <212> DNA

<213> Artificial Sequence

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<223> PCR Amplification Primer

<400> 5

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<210> 6

<211> 7

<212> PRT

<213> Artificial Sequence

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<223> Synthetic K5 Peptide

<221> VARIANT

<222> (5)...(5)

<223> Xaa = 3-I-Tyr at position 5

<400> 6

Pro Arg Lys Leu Xaa Asp Tyr
1 5

<210> 7

<211> 22

<212> DNA

<213> Artificial Sequence

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<223> Forward Primer

<400> 7

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22

<210> 8

<211> 92

<212> DNA

<213> Artificial Sequence

<220>

<223> Reverse Primer

<400> 8

attaatgaat tcctcgagcg gtccgggatc cctcggcagc ggaaccaacg gtagtgcaga
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taactggctg agcgaagaca gattgcaaag ta
92

<210> 9

<211> 111

<212> DNA

<213> Artificial Sequence

<220>

<223> Synthetic leader sequence encodes a PHO1 secretion

signal

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 60
 ttcgctcagc cagttatctg cactaccgtt ggttccgctg ccgagggatc c
 111

<210> 10
 <211> 18
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> PCR Amplification Primer

<400> 10
 gtccaggact gctaccat
 18

<210> 11
 <211> 19
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> PCR Amplification Primer

<400> 11
 ctgcttcag atgtagaga
 19

<210> 12
 <211> 2497
 <212> DNA
 <213> Homo sapiens

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 ggaagtgggtt cttctacttc ttttatttct gaaatcaggt caaggagagc ctctggatga
 120
 ctatgtgaat acccaggggg cttcactgtt cagtgtcact aagaagcagc tgggagcagg
 180
 aagtatagaa gaatgtgcag caaaatgtga ggaggacgaa gaattcacct gcagggcatt
 240
 ccaatatcac agtaaagagc aacaatgtgt gataatggct gaaaacagga agtcctccat
 300
 aatcattagg atgagagatg tagttttatt tgaaaagaaa gtgtatctct cagagtgcaa
 360
 gactgggaat ggaaagaact acagagggac gatgtccaaa acaaaaaatg gcatcacctg
 420
 tcaaaaatgg agttccactt ctccccacag acctagattc tcacctgcta cacaccctc
 480
 agagggactg gaggagaact actgcaggaa tccagacaac gatccgcagg ggccctgggtg
 540
 ctatactact gatccagaaa agagatatga ctactgcgac attcttgagt gtgaagagga
 600

atgtatgcat tgcagtggag aaaactatga cggcaaaatt tccaagacca tgtctggact
 660
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 720
 tccaaacaag aacctgaaga agaattactg tcgtaacccc gatagggagc tgcggccttg
 780
 gtgtttcacc accgacccca acaagcgctg ggaactttgt gacatccccc gctgcacaac
 840
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 900
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 960
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 1020
 ccgcaatcct gacggaaaaa gggcccccag gtgccataca accaacagcc aagtgcgggtg
 1080
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 1140
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 1200
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 1260
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 1380
 caggtgggag tactgcaacc tgaaaaaatg ctcaggaaca gaagcgagtg ttgtagcacc
 1440
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 1680
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 1860
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 1920
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 1980
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 2040
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 2100
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 2160
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 2220
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 2280
 ctgtgctggg catttgccg gaggcactga cagttgccag ggtgacagt gaggtcctct
 2340

7/13

ggtttgcttc gagaaggaca aatacatTTTt acaaggagtc acttcttggg gtcttggtg
2400
tgcacgcccc aataagcctg gtgtctatgt tcgtgtttca aggtttgtta cttggattga
2460
gggagtgatg agaaataatt aattggacgg gagacag
2497

<210> 13
<211> 23
<212> DNA
<213> Artificial Sequence

<220>
<223> PCR Amplification Primer

<400> 13
ttattaggcc gcacactgag gga
23

<210> 14
<211> 128
<212> DNA
<213> Artificial Sequence

<220>
<223> Synthetic DNA Fragment synVB1

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ccttaattaa ccgggagccc gcctaagtga cgggcttttt ttgctcttc atagtga
120
ctgacgtcg
128

<210> 15
<211> 175
<212> DNA
<213> Artificial Sequence

<220>
<223> Synthetic DNA Fragment synVB2

<400> 15
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agctgaagag ctggctcacc ttccgggtggg cctttctgcg ccttggcgcg ccaaccttaa
120
ttaaccggga gccgcgctaa tgagcgggct tttttttgct cttcacgaga cgctg
175

<210> 16
<211> 156
<212> DNA
<213> Artificial Sequence

<220>
<223> Synthetic DNA Fragment synVB3

<400> 16
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ccttcgggtg ggcctttctg cgccttggcg cgccaacctt aattaaccgg gagcccgct
120
aatgagcggg cttttttttg ctcttcacga gacgtc
156

<210> 17
<211> 172
<212> DNA
<213> Artificial Sequence

<220>
<223> Synthetic DNA Fragment synVB4

<400> 17
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tgaagagctg gctcaccttc gggtagggcct ttctgcgcct tggcgcgcca accttaatta
120
accgggagcc cgcctaata gggggctttt ttttgctctt cagagacgt cg
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<210> 18
<211> 7
<212> PRT
<213> Artificial Sequence

<220>
<223> Synthetic K5 Peptide

<221> VARIANT
<222> (7)...(7)
<223> Xaa = 3-I-Tyr at position 7

<400> 18
Pro Arg Lys Leu Tyr Asp Xaa
1 5

<210> 19
<211> 12
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA Fragment

<400> 19
catgtgaaga gc
12

<210> 20
<211> 12
<212> DNA
<213> Artificial Sequence

<220>
<223> DNA Fragment

<400> 20
gatcgctctt ca
12

<210> 21
<211> 18
<212> DNA
<213> Artificial Sequence

<220>
<223> Forward Vector Primer

<400> 21
agatctcgat cccgcgaa
18

<210> 22
<211> 18
<212> DNA
<213> Artificial Sequence

<220>
<223> Reverse Vector Primer

<400> 22
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<210> 23
<211> 21
<212> DNA
<213> Artificial Sequence

<220>
<223> Cassette Primer

<400> 23
cgggcttttt tttgctcttc a
21

<210> 24
<211> 19
<212> DNA
<213> Artificial Sequence

<220>
<223> Ubi-5p Primer

<400> 24
cagattttcg tcaagactt
19

<210> 25
<211> 18
<212> DNA

<213> Artificial Sequence

<220>

<223> Ubi-3p Primer

<400> 25

accacctctt agccttag

18

<210> 26

<211> 19

<212> DNA

<213> Artificial Sequence

<220>

<223> pET3p-ATG Primer

<400> 26

catggtatat ctccttctt

19

<210> 27

<211> 20

<212> DNA

<213> Artificial Sequence

<220>

<223> T7RevTerm Primer

<400> 27

tgagcaataa ctagcataac

20

<210> 28

<211> 18

<212> DNA

<213> Artificial Sequence

<220>

<223> pET5p Primer

<400> 28

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18

<210> 29

<211> 17

<212> DNA

<213> Artificial Sequence

<220>

<223> Strom-3p Primer

<400> 29

ttaggtctca ggggagt

17

<210> 30

<211> 19
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Strom-5p Primer

<400> 30
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 19

<210> 31
 <211> 21
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Ek-Cut-5p Primer

<400> 31
 agcggcgacg acgacgacaa g
 21

<210> 32
 <211> 21
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Ek-Cut-3p Primer

<400> 32
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 21

<210> 33
 <211> 21
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Primer

<400> 33
 tgaagagcaa aaaaaagccc g
 21

<210> 34
 <211> 101
 <212> PRT
 <213> Homo sapiens

<400> 34
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 1 5 10 15
 Glu Glu Asp Cys Met Phe Gly Asn Gly Lys Gly Tyr Arg Gly Lys Arg
 20 25 30
 Ala Thr Thr Val Thr Gly Thr Pro Cys Gln Asp Trp Ala Ala Gln Glu

```

      35              40              45
Pro His Arg His Ser Ile Phe Thr Pro Glu Thr Asn Pro Arg Ala Gly
  50              55              60
Leu Glu Lys Asn Tyr Cys Arg Asn Pro Asp Gly Asp Val Gly Gly Pro
65              70              75              80
Trp Cys Tyr Thr Thr Asn Pro Arg Lys Leu Tyr Asp Tyr Cys Asp Val
      85              90              95
Pro Gln Cys Ala Ala
      100

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<210> 35
 <211> 102
 <212> PRT
 <213> Mus musculus

```

<400> 35
Val Glu Leu Pro Thr Val Ser Gln Glu Pro Ser Gly Pro Ser Asp Ser
  1              5              10              15
Glu Thr Asp Cys Met Tyr Gly Asn Gly Lys Asp Tyr Arg Gly Lys Thr
      20              25              30
Ala Val Thr Ala Ala Gly Thr Pro Cys Gln Gly Trp Ala Ala Gln Glu
      35              40              45
Pro His Arg His Ser Ile Phe Thr Pro Gln Thr Asn Pro Arg Ala Gly
  50              55              60
Leu Glu Lys Asn Tyr Cys Arg Asn Pro Asp Gly Asp Val Asn Gly Pro
65              70              75              80
Trp Cys Tyr Thr Thr Asn Pro Arg Lys Leu Tyr Asp Tyr Cys Asp Ile
      85              90              95
Pro Leu Cys Ala Ser Ala
      100

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<210> 36
 <211> 101
 <212> PRT
 <213> Macaca mulatta

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<400> 36
Ala Ala Pro Pro Pro Val Ala Gln Leu Pro Asp Ala Glu Thr Pro Ser
  1              5              10              15
Glu Glu Asp Cys Met Phe Gly Asn Gly Lys Gly Tyr Arg Gly Lys Lys
      20              25              30
Ala Thr Thr Val Thr Gly Thr Pro Cys Gln Glu Trp Ala Ala Gln Glu
      35              40              45
Pro His Ser His Arg Ile Phe Thr Pro Glu Thr Asn Pro Arg Ala Gly
  50              55              60
Leu Glu Lys Asn Tyr Cys Arg Asn Pro Asp Gly Asp Val Gly Gly Pro
65              70              75              80
Trp Cys Tyr Thr Thr Asn Pro Arg Lys Leu Phe Asp Tyr Cys Asp Val
      85              90              95
Pro Gln Cys Ala Ala
      100

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<210> 37
 <211> 97
 <212> PRT
 <213> Bos taurus

<400> 37

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Pro Ala Ala Pro Gln Ala Pro Gly Val Glu Asn Pro Pro Glu Ala Asp
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Cys Met Ile Gly Thr Gly Lys Ser Tyr Arg Gly Lys Lys Ala Thr Thr
20
Val Ala Gly Val Pro Cys Gln Glu Trp Ala Ala Gln Glu Pro His His
35      40      45
His Ser Ile Phe Thr Pro Glu Thr Asn Pro Gln Ser Gly Leu Glu Arg
50      55      60
Asn Tyr Cys Arg Asn Pro Asp Gly Asp Val Asn Gly Pro Trp Cys Tyr
65      70      75      80
Thr Met Asn Pro Arg Lys Leu Phe Asp Tyr Cys Asp Val Pro Gln Cys
85      90      95
Glu

```

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<210> 38
<211> 100
<212> PRT
<213> Sus scrofa

```

```

<400> 38
Thr Asn Phe Pro Ala Ile Ala Gln Val Pro Ser Val Glu Asp Leu Ser
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Glu Asp Cys Met Phe Gly Asn Gly Lys Arg Tyr Arg Gly Lys Arg Ala
20      25      30
Thr Thr Val Ala Gly Val Pro Cys Gln Glu Trp Ala Ala Gln Glu Pro
35      40      45
His Arg His Ser Ile Phe Thr Pro Glu Thr Asn Pro Arg Ala Gly Leu
50      55      60
Glu Lys Asn Tyr Cys Arg Asn Pro Asp Gly Asp Asp Asn Gly Pro Trp
65      70      75      80
Cys Tyr Thr Thr Asn Pro Gln Lys Leu Phe Asp Tyr Cys Asp Val Pro
85      90      95
Gln Cys Val Thr
100

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<210> 39
<211> 7
<212> PRT
<213> Homo sapiens

```

```

<400> 39
Pro Glu Lys Arg Tyr Asp Tyr
1      5

```

```

<210> 40
<211> 31
<212> PRT
<213> Homo sapiens

```

```

<400> 40
Gln Asp Trp Ala Ala Gln Glu Pro His Arg His Ser Ile Phe Thr Pro
1      5      10      15
Glu Thr Pro Glu Thr Asn Pro Arg Ala Gly Leu Glu Lys Asn Tyr
20      25      30

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